

Linking CoCoMac and Brede databases

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Linking CoCoMac and Brede databases

CoCoMac database records anatomical connectivity of the macaque.

Brede database contains stereotaxic coordinates in the human.

Combining these databases will enable visualization of the 3-dimensional connectivity.

Project with Jesper Rønager, Copenhagen University Hospital Rigshospitalet, Neurology.

Some functionality already exists via the Catacomb and Carat software (Kötter, 2004; Cannon et al., 2003; Van Essen et al., 2001), e.g., a network with 95 nodes and 2402 connections has been constructed (Kaiser and Hilgetag, 2004; Sporns et al., 2004).

CoCoMac connectivity database

Connectivity output list, PrimaryProjections

34 Items, page 1/2 select page: [1](#) [2](#)

SearchString: ('CD') [KEYWORDS]

details

output type: [HTML -> Brow](#)

items per page: [20](#)

order by: [SourceMap](#)

[ascending](#)

☐ user comments [display](#)

[display all results](#) [edit search](#) [show url](#) [back to search](#) [start new search](#)

Item	SourceSite	PDC	Hemisph.	Density	PDC	Course	TargetSite	PDC	Hemisph.	Laminae
1. <input type="checkbox"/>	B09-19	D	?	X	-	I	BD77-Cd	A	?	Laminae LS
2. <input type="checkbox"/>	B09-19	D	?	X	-	I	BD77-Cd	A	?	Laminae LS
3. <input type="checkbox"/>	B09-18	D	?	X	-	I	BD77-Cd	A	?	Laminae LS
4. <input type="checkbox"/>	B09-18	D	?	X	-	I	BD77-Cd	A	?	Laminae LS
5. <input type="checkbox"/>	B09-18	D	?	X	-	I	BD77-Cd	A	?	Laminae LS
6. <input type="checkbox"/>	B09-18	D	?	X	-	?	BD77-Cd	F	?	Laminae LS
7. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdr	C	L	Laminae LS
8. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdr	C	L	Laminae LS
9. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-46sup	C	L	Laminae LS
10. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdc	C	L	Laminae LS
11. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-46inf	C	L	Laminae LS
12. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMvc	C	L	Laminae LS
13. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMv	C	L	Laminae LS
14. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdc	C	L	Laminae LS
15. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdc	C	L	Laminae LS
16. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMdc	C	L	Laminae LS
17. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-SMA	C	L	Laminae LS
18. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-PMr	C	L	Laminae LS
19. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-SMA	C	L	Laminae LS
20. <input type="checkbox"/>	RTMB99-Cd	L	L	0	-	I	RTMB99-M1	C	L	Laminae LS

CoCoMac records anatomical connectivity in the Macaque brain with data from presently 410 papers (Stephan et al., 2001).

Brain region ontology (Stephan et al., 2000).

Stores “from”, “to” and how strong the link is, what tracer, etc.

Database on the Internet with output as HTML or XML.

CoCoMac brain region ontology (“mapping”)

An entry in the “mapping” table of the CoCoMac database records:

“AI92-23 I AP84-23 AI92 Figs. 2-3”

This means:

“Area 23 of (Amaral and Insausti, 1992) is identical to area 23 of (Amaral and Price, 1984) according to (Amaral and Insausti, 1992, Figs. 2–3)”

Other “23” areas:

AI92-23, AP84-23, ASM94-23, B05-23, B09-23, B88-23, . . . , V93-23, VPR87-23, YP88-23

CoCoMac brain region ontology (“mapping”)

CoCoMac brain regions (brain sites) are often ordered in a hierarchy, e.g.:

B09-23 L PHT00-23a PHT00

... means Brodmann area 23 is larger than (a suprastructure of) 23a of (Paxinos et al., 1999) (!?).

Particularly the so-called generalized hierarchy names (GM ontology) (Kötter and Wanke, 2005).

Connectivity in CoCoMac

Connectivity entry:

AI92-23 A L 0 - I AI92-Bi A L

means

Area 23 of (Amaral and Insausti, 1992) in the left (L) hemisphere has zero connection (0) to “intermediate part of the basal amygdaloid nucleus” (Bi) of (Amaral and Insausti, 1992) in the left (L) hemisphere. Both areas are named explicitly (A).

Both mapping and connectivity are available as XML

Brede Database

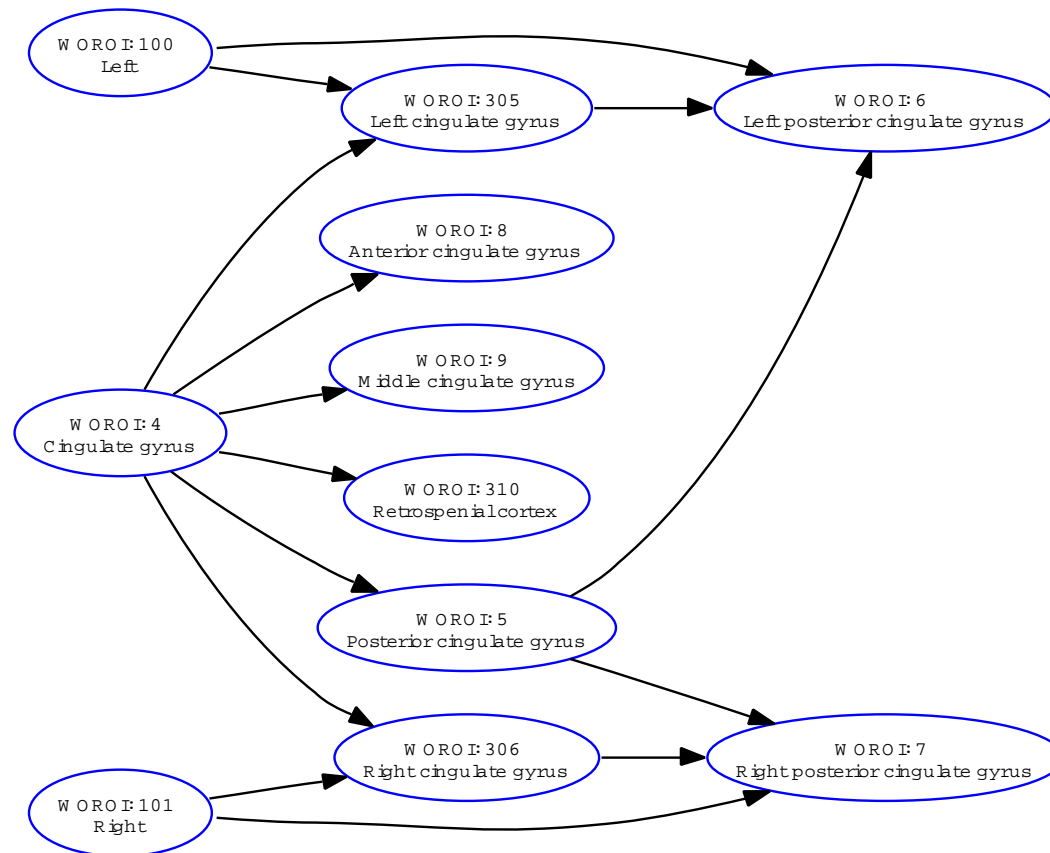
x	y	z	Lobar anatomy	Functional area
-50	-2	36		Left frontal eye field
42	-8	48		Right frontal eye field
4	-10	60		Left and right supplementary eye field
-20	-14	4	Left putamen and thalamus	
22	-8	8	Right putamen	
20	-20	16	Right thalamus	
16	-64	-20	Vermis	
-20	-54	-32	Flocculus region	
-6	-38	-4	Mesencephalon	
-18	28	-20		Left orbit (eye muscles)
20	28	-20		Right orbit (eye muscles)
-12	-86	32	Left cuneus	
8	-86	32	Right cuneus	

Brede Database (Nielsen, 2003).

Interesting fields here: 3906 “location” structures with 3D stereotaxic coordinates, lobar anatomy and functional area textual labels and Brodmann areas.

Functions in the Brede Toolbox (Nielsen and Hansen, 2000) are able to find all coordinates with a given lobar anatomy label.

Brede brain region taxonomy



Organizes brain areas in a hierarchy.

Variations on naming of a brain area.

Extended significantly to link to the detailed brain areas from CoCoMac.

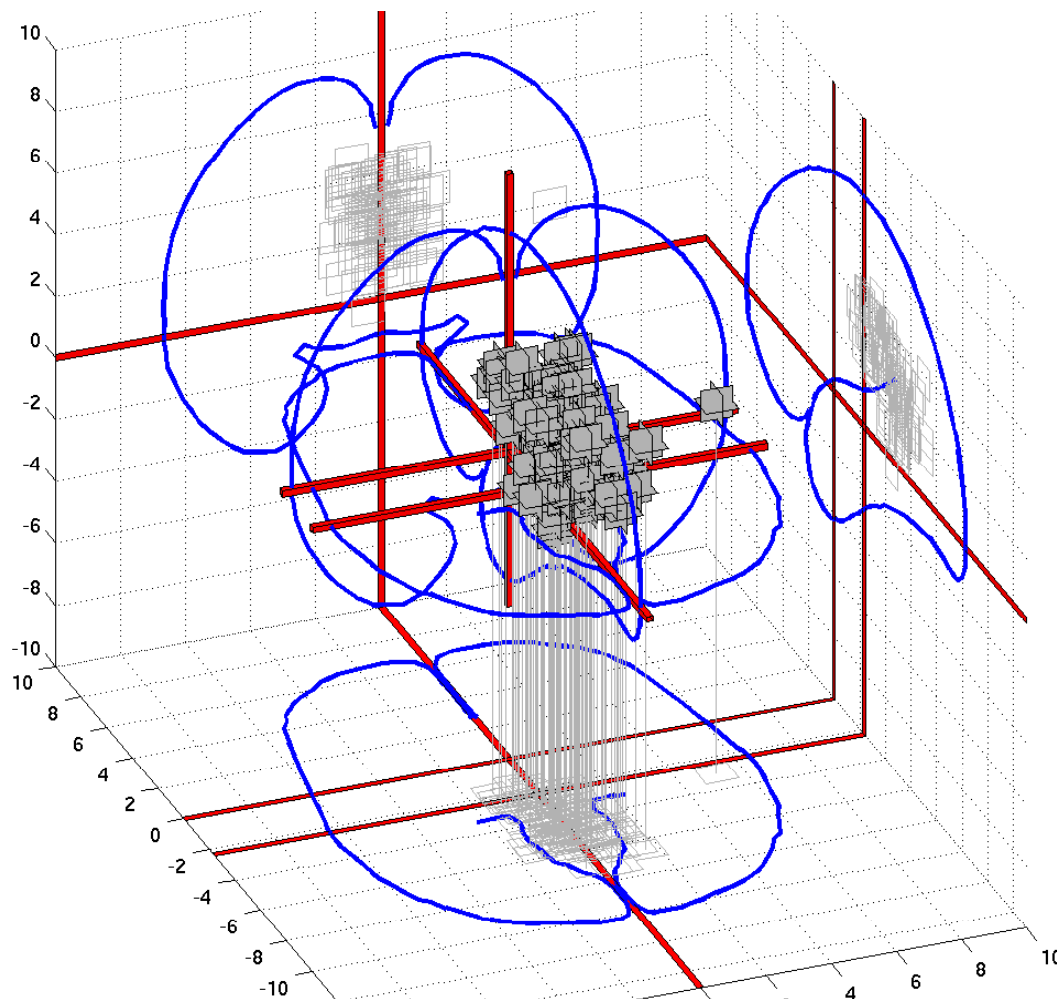
Links to stereotaxic volumetric definitions of brain area (Tzourio-Mazoyer et al., 2002; Hammers et al., 2002).

Figure 1: Brede brain region taxonomy at cingulate gyrus.

Example entry in XML of the Brede Database

```
<Roi>
  <woroi>5</woroi>
  <name>Posterior cingulate gyrus</name>
  <abbreviation>PCgG</abbreviation>
  <abbreviation>CGp</abbreviation>
  <brainInfo>144</brainInfo>
  <cocomacSite>OMG96-CGp</cocomacSite>
  <type>roi</type>
  <variation>Posterior cingulate</variation>
  <variation>Posterior cingulate area</variation>
  <variation>Posterior gyrus cinguli</variation>
  <variation>Posterior cingulate cortex</variation>
  <parent>4</parent>
</Roi>
```

Finding a representative coordinate



Search on lobar anatomy, if no coordinates are found try parent (supra-region).

Problem with, e.g., 8a and 8b which fall back on coordinates labeled BA8.

Model the distribution of stereotaxic coordinates with kernel density modeling (Nielsen and Hansen, 2002) and pick the coordinate with the highest probability density.

Matching brain areas in CoCoMac and Brede

Explicitly (manually) added individual CoCoMac brain sites to their specific entry in the Brede brain region taxonomy.

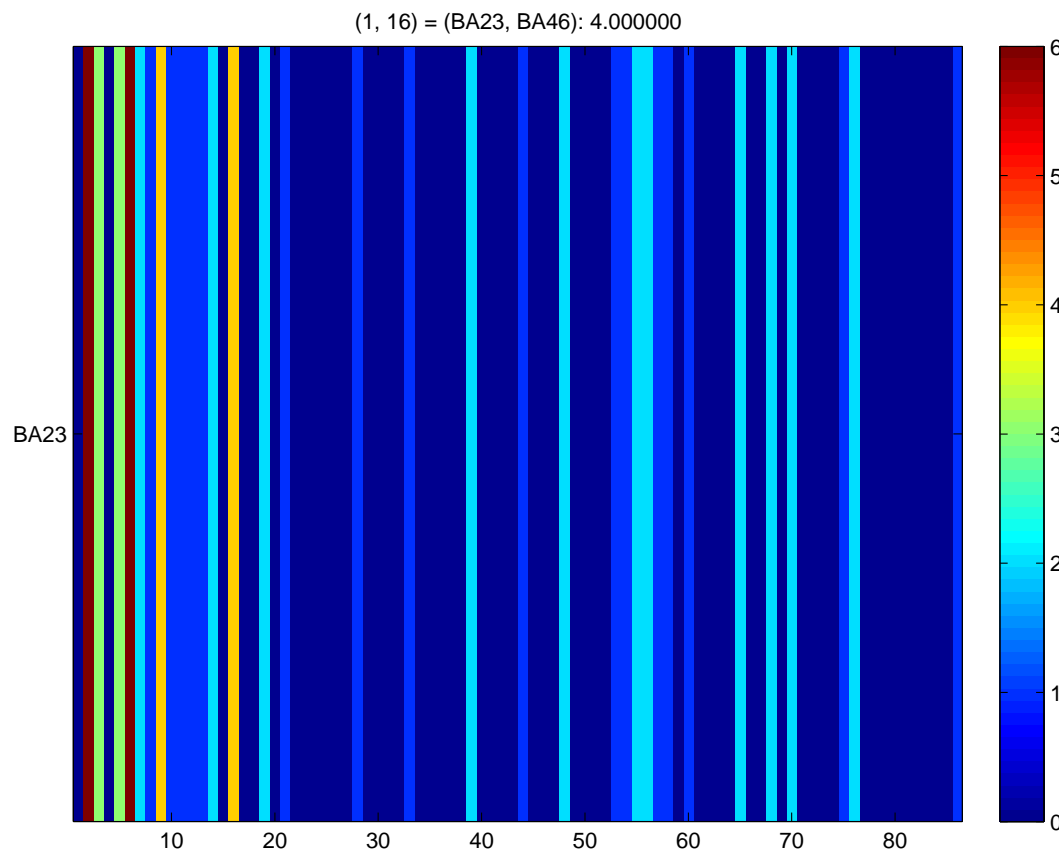
Helped by NeuroNames (Bowden and Martin, 1995), atlases (Mai et al., 1997) and texts with human/macaque comparative studies, e.g., (Van Essen, 2003; Scott and Johnsrude, 2003).

What to do about certain area (macaque and human brains are not completely homologous), e.g., Brodmann areas 13, 14, 15 and 16 are defined for monkeys — not humans?

Examples on presently missing matches:

"PBK86-region 1", VV19-4a, RACR99-V1_V (i.e., layer five of visual area 1), SA94b-ECL (Caudal limiting field of entorhinal cortex), PK85-1_Face (i.e., face area of Brodmann area 1), ... and 625 others.

Example on connectivity matrix



308 entries for area 23 (i.e., BA23) as source brain site when querying CoCoMac.

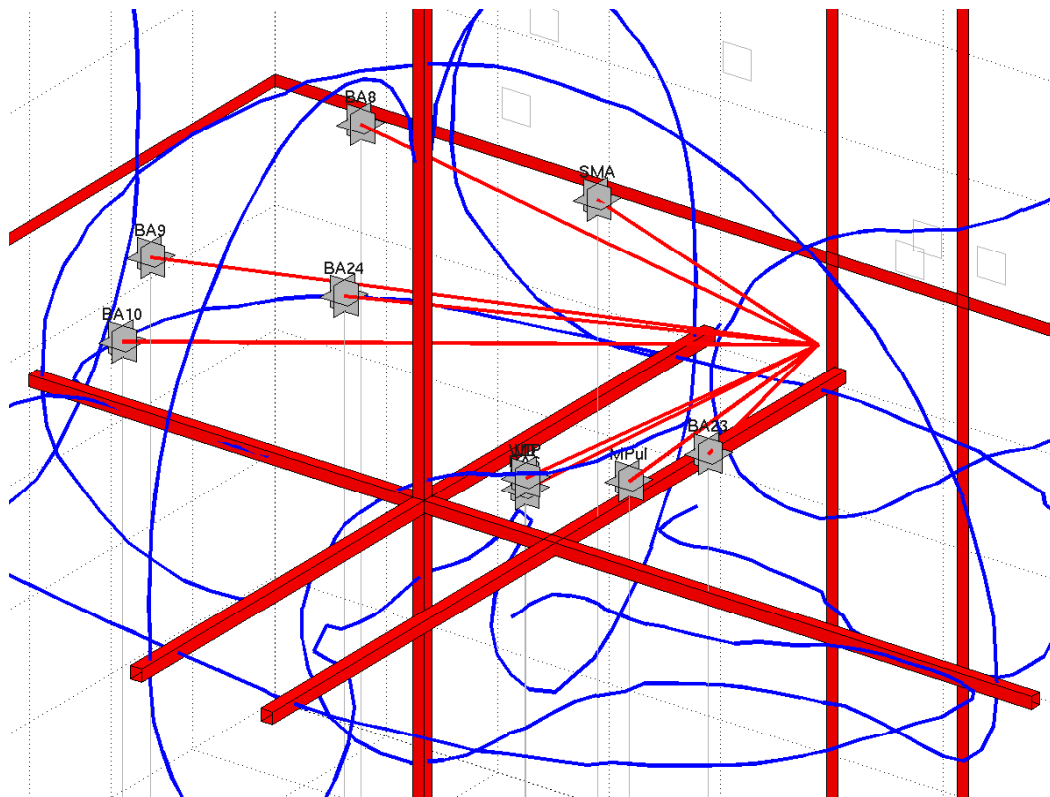
27 unmatched to Brede brain region taxonomy.

86 brain areas left.

33 brain areas with non-zero connections

Figure 2: Connection-"matrix" from BA23.

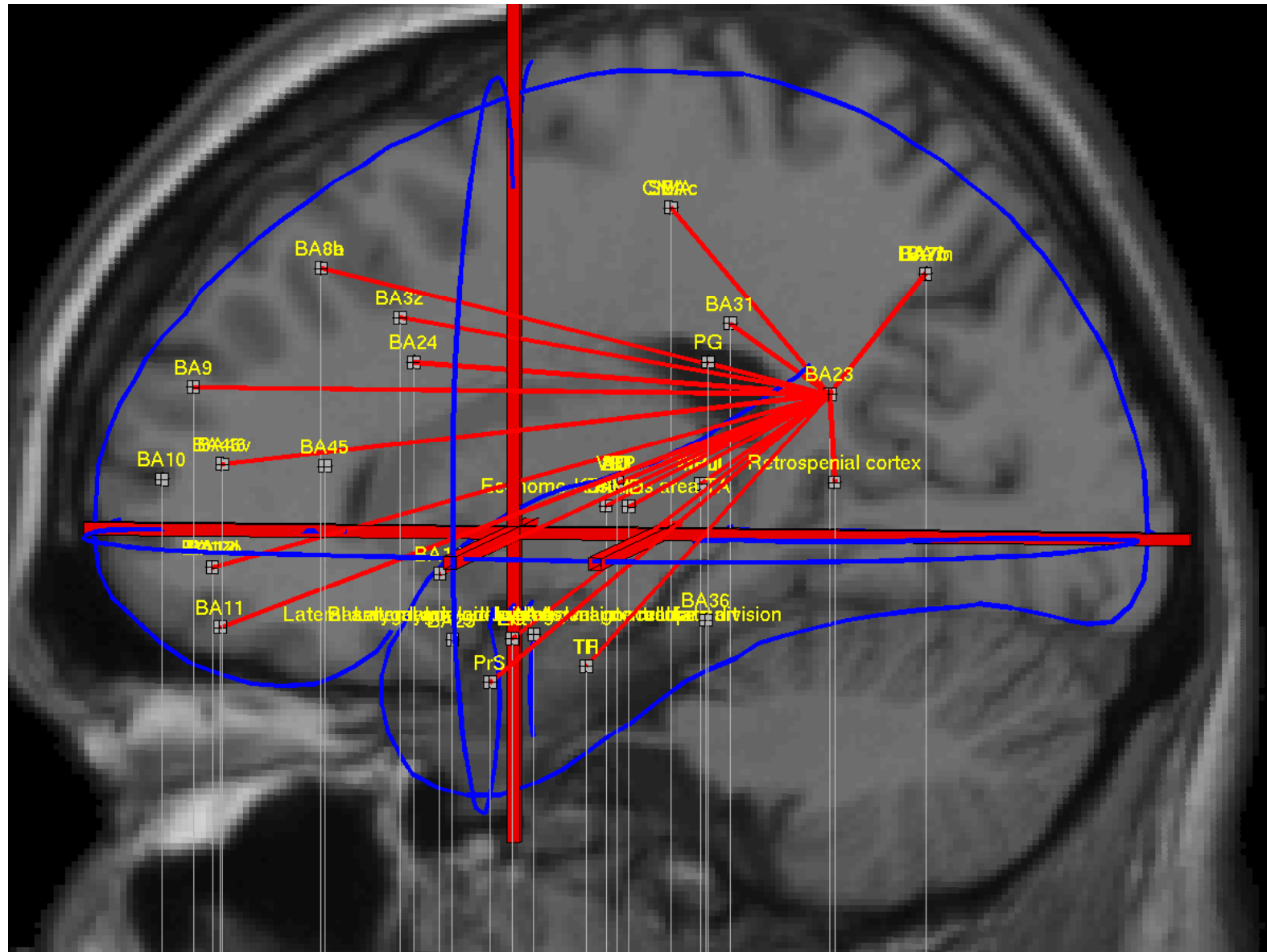
Example 3D visualization



Query CoCoMac database for connections *from* BA7 (pre-cuneus).

Here no distinction between left and right.

Figure 3: Connections from BA7. 3D plot from left posterior.



Summary

Brain region taxonomy in the Brede Database originally developed for human molecular neuroimaging extended to accomodate (many) brain sites of CoCoMac.

Brede Toolbox extended to handle CoCoMac data and match it against information in the Brede Database.

It is difficult/problematic to match all brain areas.

The major part of macaque connections in CoCoMac can be plotted in 3D human stereotaxic space.

References

- Amaral, D. G. and Insausti, R. (1992). Retrograde transport of D-[³H]-aspartate into the monkey amygdaloid complex. *Experimental Brain Research*, 88(2):375–388. DOI: 10.1007/BF02259113.
- Amaral, D. G. and Price, J. L. (1984). Amygdalo-cortical projections in the monkey (*Macaca fascicularis*). *Journal of Comparative Neurology*, 230(4). PMID: 6520247.
- Bowden, D. M. and Martin, R. F. (1995). NeuroNames brain hierarchy. *NeuroImage*, 2(1):63–84. PMID: 9410576. ISSN 1053-8119.
- Cannon, R. C., Hasselmo, M. E., and Koene, R. A. (2003). From biophysics to behavior. catacomb2 and the design of biologically-plausible models for spatial navigation. *Neuroinformatics*, 1(1):3–42. PMID: 15055391. <http://www.anc.ed.ac.uk/~cannon/catacomb2.pdf>.
- Hammers, A., Koepp, M. J., Free, S. L., Brett, M., Richardson, M. P., Labbé, C., Cunningham, V. J., Brooks, D. J., and Duncan, J. (2002). Implementation and application of a brain template for multiple volumes of interest. *Human Brain Mapping*, 15(3):165–174. DOI: 10.1002/hbm.10016. <http://www3.interscience.wiley.com/cgi-bin/abstract/89013541/>. ISSN 1065-9471. Describes a segmentation of the MNI single subject brain. Assessment of the method by using manual labeling of landmarks and exemplified on a FMZ PET study.
- Kaiser, M. and Hilgetag, C. C. (2004). Modelling the development of cortical systems networks. *Neurocomputing*, 58–60:297–302. DOI: 10.1016/j.neucom.2004.01.059. <http://www.biological-networks.org/pubs/Kaiser2004c.pdf>.
- Kötter, R. (2004). Online retrieval, processing, and visualization of primate connectivity data from the CoCoMac database. *Neuroinformatics*, 2(2):127–144. PMID: 15319511. <http://www.cocomac.org/cocomac2004.pdf>.
- Kötter, R. and Wanke, E. (2005). Mapping brains without coordinates. *Philosophical Transactions of the Royal Society, Series B, Biological Sciences*, 360(1456):751–766. DOI: 10.1098.rstb.2005.1625.

- Mai, J. K., Assheuer, J., and Paxinos, G. (1997). *Atlas of the Human Brain*. Academic Press, San Diego, California. ISBN 0124653618.
- Nielsen, F. Å. (2003). The Brede database: a small database for functional neuroimaging. *NeuroImage*, 19(2). <http://208.164.121.55/hbm2003/abstract/abstract906.htm>. Presented at the 9th International Conference on Functional Mapping of the Human Brain, June 19–22, 2003, New York, NY. Available on CD-Rom.
- Nielsen, F. Å. and Hansen, L. K. (2000). Experiences with Matlab and VRML in functional neuroimaging visualizations. In Klasky, S. and Thorpe, S., editors, *VDE2000 - Visualization Development Environments, Workshop Proceedings, Princeton, New Jersey, USA, April 27–28, 2000*, pages 76–81, Princeton, New Jersey. Princeton Plasma Physics Laboratory. http://www.imm.dtu.dk/pubdb/views/edoc_download.php/1231/pdf/imm1231.pdf. CiteSeer: <http://citeseer.ist.psu.edu/309470.html>.
- Nielsen, F. Å. and Hansen, L. K. (2002). Modeling of activation data in the BrainMap™ database: Detection of outliers. *Human Brain Mapping*, 15(3):146–156. DOI: 10.1002/hbm.10012. <http://www3.interscience.wiley.com/cgi-bin/abstract/89013001/>. CiteSeer: <http://citeseer.ist.psu.edu/nielsen02modeling.html>.
- Paxinos, G., Huang, X.-F., and Toga, A. W. (1999). *The Rhesus monkey brain in stereotaxic coordinates*. Academic Press. ISBN 0123582555.
- Scott, S. K. and Johnsrude, I. S. (2003). The neuroanatomical and functional organization of speech perception. *Trends in Neurosciences*, 26(2):100–107. DOI: 10.1016/S0166-2236(02)00037-1. http://www-visl.technion.ac.il/karniel/CMCC/Neuroanatomy_of_speech_perception.pdf.
- Sporns, O., Chialvo, D. R., Kaiser, M., and Hilgetag, C. C. (2004). Organization, development and function of complex brain networks. *Trends in Cognitive Sciences*, 8(9):418–425.
- Stephan, K. E., Kamper, L., Bozkurt, A., Burns, G. A. P. C., Young, M. P., and Kötter, R. (2001). Advanced database methodology for the collation of connectivity data on the macaque brain (Co-CoMac). *Philosophical Transactions for the Royal Society, London, Series B, Biological Sciences*, 356(1412):1159–1186. PMID: 11545697. http://www.cocomac.org/cocomac_paper.pdf.

- Stephan, K. E., Zilles, K., and Kötter, R. (2000). Coordinate-independent mapping of structural and functional data by objective relational transformation (ORT). *Philosophical Transactions of the Royal Society, London, Series B, Biological Sciences*, 355(1393):37–54. PMID: 10703043.
- Tzourio-Mazoyer, N., Landeau, B., Papathanassiou, D., Crivello, F., Etard, O., Delcroix, N., Mazoyer, B., and Joliot, M. (2002). Automated anatomical labeling of activations in SPM using a macroscopic anatomical parcellation of the MNI MRI single-subject brain. *NeuroImage*, 15(1):273–289. DOI: 10.1006/nimg.2001.0978.
- Van Essen, D. C. (2003). Organization of visual areas in macaque and human cerebral cortex. In Chalupa, L. M. and Werner, J. S., editors, *The Visual Neurosciences*, chapter 32, pages 507–521. MIT Press, Cambridge, MA. <http://brainmap.wustl.edu/resources/VisChapter.pdf>. ISBN 0262033089.
- Van Essen, D. C., Drury, H. A., Dickson, J., Harwell, J., Hanlon, D., , and Anderson, C. H. (2001). An integrated software suite for surface-based analyses of cerebral cortex. *Journal of the American Medical Informatics Association*, 8(5):443–459. PMID: 11522765.